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Double-blind peer review submissions: write DBPR and your manuscript number here instead of author names.

Last updated by author(s): YYYY-MM-DD

Reporting Summary

Nature Research wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Research policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

Sta	itistics					
For	all statistical an	alyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.				
n/a	Confirmed					
	The exact	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement				
	A stateme	ent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly				
	The statistical test(s) used AND whether they are one- or two-sided Only common tests should be described solely by name; describe more complex techniques in the Methods section.					
\boxtimes	A descript	ion of all covariates tested				
	A descript	ion of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons				
	A full description of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)					
	For null hypothesis testing, the test statistic (e.g. <i>F</i> , <i>t</i> , <i>r</i>) with confidence intervals, effect sizes, degrees of freedom and <i>P</i> value noted Give <i>P</i> values as exact values whenever suitable.					
\boxtimes	For Bayes	ian analysis, information on the choice of priors and Markov chain Monte Carlo settings				
	For hierar	chical and complex designs, identification of the appropriate level for tests and full reporting of outcomes				
\boxtimes	\square Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated					
		Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.				
So	ftware an	d code				
Poli	cy information	about <u>availability of computer code</u>				
Da	ata collection	The code of the project after the manuscript acceptance will be released open source.				
Da	nta analysis	Please refer to the rebuttal letter, in short after acceptance all codes will be shared open source.				
		custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors and encourage code deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.				

Data

Policy information about <u>availability of data</u>

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A list of figures that have associated raw data $% \left(1\right) =\left(1\right) \left(1\right) \left($
- A description of any restrictions on data availability

No patient data will be shared to the reader. However, some patient data in python format has been shared with the reviewer.

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Please select the o	ne below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.						
∑ Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences						
For a reference copy of	the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>						
Life scier	nces study design						
All studies must dis	sclose on these points even when the disclosure is negative.						
Sample size	Describe how sample size was determined, detailing any statistical methods used to predetermine sample size OR if no sample-size calculation was performed, describe how sample sizes were chosen and provide a rationale for why these sample sizes are sufficient.						
Data exclusions	Describe any data exclusions. If no data were excluded from the analyses, state so OR if data were excluded, describe the exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.						
Replication	Describe the measures taken to verify the reproducibility of the experimental findings. If all attempts at replication were successful, confirm this OR if there are any findings that were not replicated or cannot be reproduced, note this and describe why.						
Randomization	Describe how samples/organisms/participants were allocated into experimental groups. If allocation was not random, describe how covariates were controlled OR if this is not relevant to your study, explain why.						
Blinding	Describe whether the investigators were blinded to group allocation during data collection and/or analysis. If blinding was not possible, describe why OR explain why blinding was not relevant to your study.						

Reporting for specific materials, systems and methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, system or method listed is relevant to your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.

Materials & experimental systems			Methods				
n/a	Involved in the study	n/a	Involved in the study				
	Antibodies		ChIP-seq				
	Eukaryotic cell lines		Flow cytometry				
	Palaeontology and archaeology		MRI-based neuroimaging				
	Animals and other organisms						
	Human research participants						
	Clinical data						
	Dual use research of concern						

Antibodies

Antibodies used

Describe all antibodies used in the study; as applicable, provide supplier name, catalog number, clone name, and lot number.

Validation

Describe the validation of each primary antibody for the species and application, noting any validation statements on the manufacturer's website, relevant citations, antibody profiles in online databases, or data provided in the manuscript.

Eukaryotic cell lines

Policy information about cell lines Cell line source(s) State the source of each cell line used. Authentication Describe the authentication procedures for each cell line used OR declare that none of the cell lines used were authenticated. Mycoplasma contamination Confirm that all cell lines tested negative for mycoplasma contamination OR describe the results of the testing for mycoplasma contamination OR declare that the cell lines were not tested for mycoplasma contamination. Commonly misidentified lines Name any commonly misidentified cell lines used in the study and provide a rationale for their use. (See ICLAC register)

Palaeontology and Archaeology

Specimen provenance

Provide provenance information for specimens and describe permits that were obtained for the work (including the name of the issuing authority, the date of issue, and any identifying information).

Specimen deposition

Indicate where the specimens have been deposited to permit free access by other researchers.

Dating methods

If new dates are provided, describe how they were obtained (e.g. collection, storage, sample pretreatment and measurement), where they were obtained (i.e. lab name), the calibration program and the protocol for quality assurance OR state that no new dates are provided.

Tick this box to confirm that the raw and calibrated dates are available in the paper or in Supplementary Information.

Ethics oversight

Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or guidance was required and explain why not.

Note that full information on the approval of the study protocol must also be provided in the manuscript.

Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

Laboratory animals

For laboratory animals, report species, strain, sex and age OR state that the study did not involve laboratory animals.

Wild animals

Provide details on animals observed in or captured in the field; report species, sex and age where possible. Describe how animals were caught and transported and what happened to captive animals after the study (if killed, explain why and describe method; if released, say where and when) OR state that the study did not involve wild animals.

Field-collected samples

For laboratory work with field-collected samples, describe all relevant parameters such as housing, maintenance, temperature, photoperiod and end-of-experiment protocol OR state that the study did not involve samples collected from the field.

Ethics oversight

Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or guidance was required and explain why not.

Note that full information on the approval of the study protocol must also be provided in the manuscript.

Human research participants

Policy information about studies involving human research participants

Population characteristics

Describe the covariate-relevant population characteristics of the human research participants (e.g. age, gender, genotypic information, past and current diagnosis and treatment categories). If you filled out the behavioural & social sciences study design questions and have nothing to add here, write "See above."

Recruitment

Describe how participants were recruited. Outline any potential self-selection bias or other biases that may be present and how these are likely to impact results.

Ethics oversight

Identify the organization(s) that approved the study protocol.

Note that full information on the approval of the study protocol must also be provided in the manuscript.

Clinical data

Policy information about <u>clinical studies</u>

All manuscripts should comply with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration

Provide the trial registration number from ClinicalTrials.gov or an equivalent agency.

Study protocol

Note where the full trial protocol can be accessed OR if not available, explain why.

Data collection

Describe the settings and locales of data collection, noting the time periods of recruitment and data collection.

Outcomes

Describe how you pre-defined primary and secondary outcome measures and how you assessed these measures.

Dual use research of concern

Policy information about <u>dual use research of concern</u>

Hazards

Could the accidental, deli in the manuscript, pose a	perate or reckless misuse of agents or technologies generated in the work, or the application of information presented threat to:					
No Yes						
Public health						
National security						
Crops and/or livest	ock					
Ecosystems						
Any other significant area						
Experiments of concer	n					
Does the work involve an	y of these experiments of concern:					
No Yes						
Demonstrate how	to render a vaccine ineffective					
Confer resistance t	o therapeutically useful antibiotics or antiviral agents					
Enhance the virule	nce of a pathogen or render a nonpathogen virulent					
Increase transmissi	bility of a pathogen					
Alter the host rang	e of a pathogen					
Enable evasion of o	liagnostic/detection modalities					
Enable the weapor	ization of a biological agent or toxin					
Any other potentia	lly harmful combination of experiments and agents					
ChID cog						
ChIP-seq						
Data deposition						
Confirm that both raw	and final processed data have been deposited in a public database such as GEO.					
Confirm that you have	deposited or provided access to graph files (e.g. BED files) for the called peaks.					
Data access links	For "Initial submission" or "Revised version" documents, provide reviewer access links. For your "Final submission" document,					
May remain private before public	ation. provide a link to the deposited data.					
Files in database submiss	on Provide a list of all files available in the database submission.					
Genome browser session (e.g. <u>UCSC</u>)	Provide a link to an anonymized genome browser session for "Initial submission" and "Revised version" documents only, to enable peer review. Write "no longer applicable" for "Final submission" documents.					
Methodology						
Replicates	Describe the experimental replicates, specifying number, type and replicate agreement.					
Sequencing depth	Describe the sequencing depth for each experiment, providing the total number of reads, uniquely mapped reads, length of reads and whether they were paired- or single-end.					
Antibodies	Describe the antibodies used for the ChIP-seq experiments; as applicable, provide supplier name, catalog number, clone name, and lot number.					
Peak calling parameters	Specify the command line program and parameters used for read mapping and peak calling, including the ChIP, control and index files used.					
Data quality	escribe the methods used to ensure data quality in full detail, including how many peaks are at FDR 5% and above 5-fold enrichment.					
Software	Describe the software used to collect and analyze the ChIP-seq data. For custom code that has been deposited into a community					

Flow Cytometry						
Plots Confirm that:						
	and fluorochrome used (e.g. CD4-FITC).					
	. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).					
All plots are contour plots with c	outliers or pseudocolor plots.					
A numerical value for number of	f cells or percentage (with statistics) is provided.					
Methodology						
Sample preparation De	scribe the sample preparation, detailing the biological source of the cells and any tissue processing steps used.					
Instrument	entify the instrument used for data collection, specifying make and model number.					
	scribe the software used to collect and analyze the flow cytometry data. For custom code that has been deposited into a mmunity repository, provide accession details.					
	scribe the abundance of the relevant cell populations within post-sort fractions, providing details on the purity of the mples and how it was determined.					
0 07	ribe the gating strategy used for all relevant experiments, specifying the preliminary FSC/SSC gates of the starting cell ulation, indicating where boundaries between "positive" and "negative" staining cell populations are defined.					
Tick this box to confirm that a fig	gure exemplifying the gating strategy is provided in the Supplementary Information.					
Magnetic resonance ima	aging					
Experimental design						
Design type	Indicate task or resting state; event-related or block design.					
Design specifications	Specify the number of blocks, trials or experimental units per session and/or subject, and specify the length of each trial or block (if trials are blocked) and interval between trials.					
Behavioral performance measures	State number and/or type of variables recorded (e.g. correct button press, response time) and what statistics were used to establish that the subjects were performing the task as expected (e.g. mean, range, and/or standard deviation across subjects).					
Acquisition						
Imaging type(s)	Specify: functional, structural, diffusion, perfusion.					
Field strength	Specify in Tesla					
Sequence & imaging parameters	Specify the pulse sequence type (gradient echo, spin echo, etc.), imaging type (EPI, spiral, etc.), field of view, matrix size, slice thickness, orientation and TE/TR/flip angle.					
Area of acquisition	State whether a whole brain scan was used OR define the area of acquisition, describing how the region was determined.					
Diffusion MRI Used	☐ Not used					

Preprocessing

Normalization

Normalization template

Noise and artifact removal

Preprocessing software Provide detail on software version and revision number and on specific parameters (model/functions, brain extraction, segmentation, smoothing kernel size, etc.).

If data were normalized/standardized, describe the approach(es): specify linear or non-linear and define image types used for transformation OR indicate that data were not normalized and explain rationale for lack of normalization.

Describe the template used for normalization/transformation, specifying subject space or group standardized space (e.g. original Talairach, MNI305, ICBM152) OR indicate that the data were not normalized.

Congress of the second second

Describe your procedure(s) for artifact and structured noise removal, specifying motion parameters, tissue signals and physiological signals (heart rate, respiration).

Volume censoring

Define your software and/or method and criteria for volume censoring, and state the extent of such censoring.

Statistica	I mode	ling	&	infer	ence
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datistical modeling & intere	nec .						
Model type and settings	Specify type (mass univariate, multivariate, RSA, predictive, etc.) and describe essential details of the model at the first and second levels (e.g. fixed, random or mixed effects; drift or auto-correlation).						
Effect(s) tested	Define precise effect in terms of the task or stimulus conditions instead of psychological concepts and indicate whether ANOVA or factorial designs were used.						
Specify type of analysis: W	hole brain ROI-based Both						
Statistic type for inference (See Eklund et al. 2016)	Specify voxel-wise or cluster-wise and report all relevant parameters for cluster-wise methods.						
Correction	Describe the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte Carlo).						
Models & analysis n/a Involved in the study							
Functional and/or effective conn	Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation, mutual information).						
Graph analysis	Report the dependent variable and connectivity measure, specifying weighted graph or binarized graph, subject- or group-level, and the global and/or node summaries used (e.g. clustering coefficient, efficiency, etc.).						

Multivariate modeling and predictive analysis

Specify independent variables, features extraction and dimension reduction, model, training and evaluation metrics.